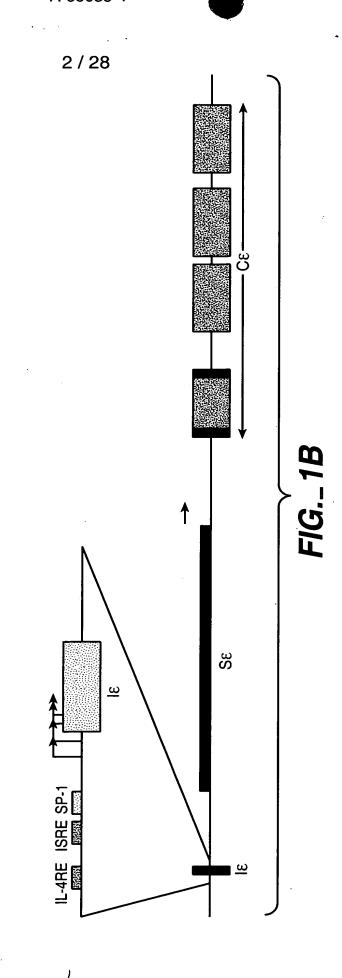
FIG._1A

- 1 GCTGGGCTAA ACTGGGCTAG CCTGAGCTGG GCTGAACTGG GCTGCTGGGC
- 51 TGGACTGGGT AAGCTGGGCT GAGCTGGGTT GGGTGGAAAT GGGCTGAGCT

FIG._2B

- 1 GGTTTGGCTG GGCTGGGCTG GGCTGGGCTG GGTTCAGCTG AGCGGGTTGG
- 51 GTTAGACTGG GTCAAACTGG TTCAGC

FIG._2C



GERMLINE & LOCUS

IL-4 INDUCIBLE PROMOTER FRAGMENT

IL-4RE ISRE SP-1

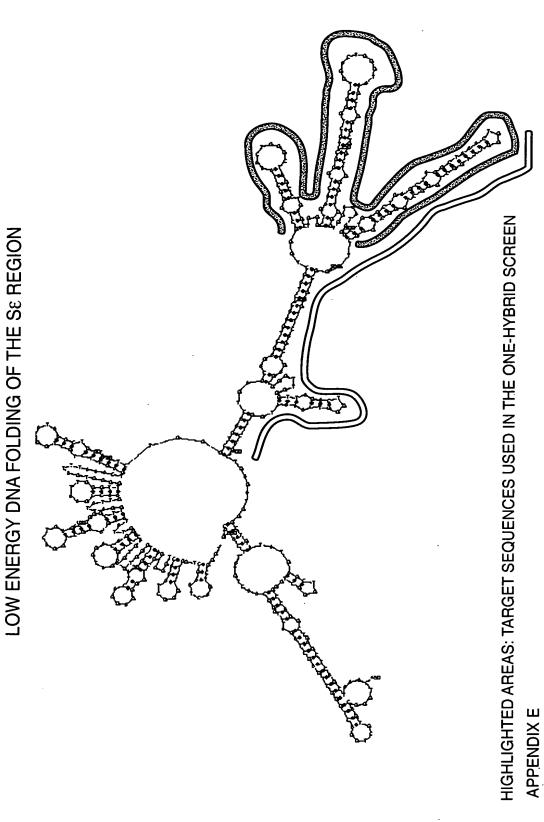


FIG._2A

+

YEAST ONE-HYBRID SCREENING

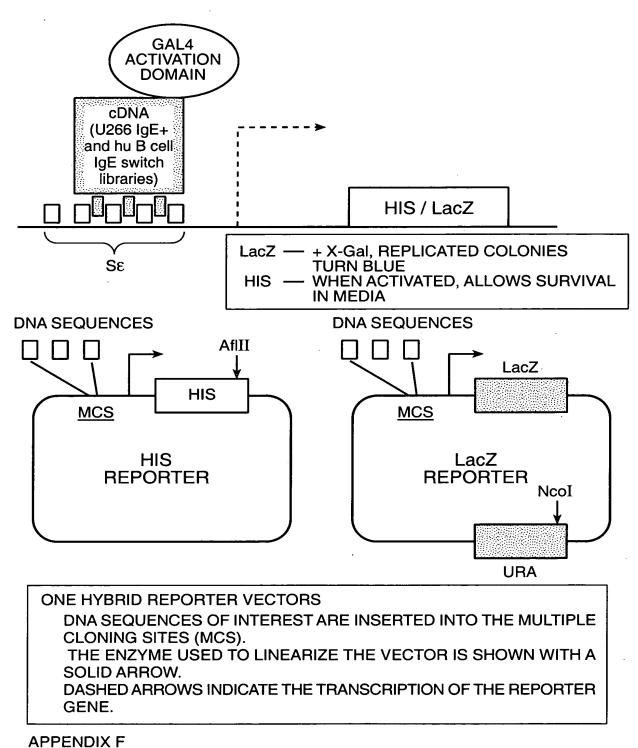
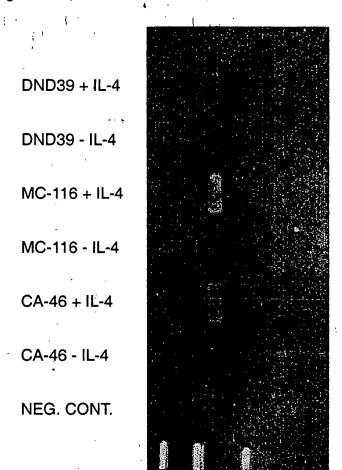


FIG._3

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BEST AVAILABLE COPY

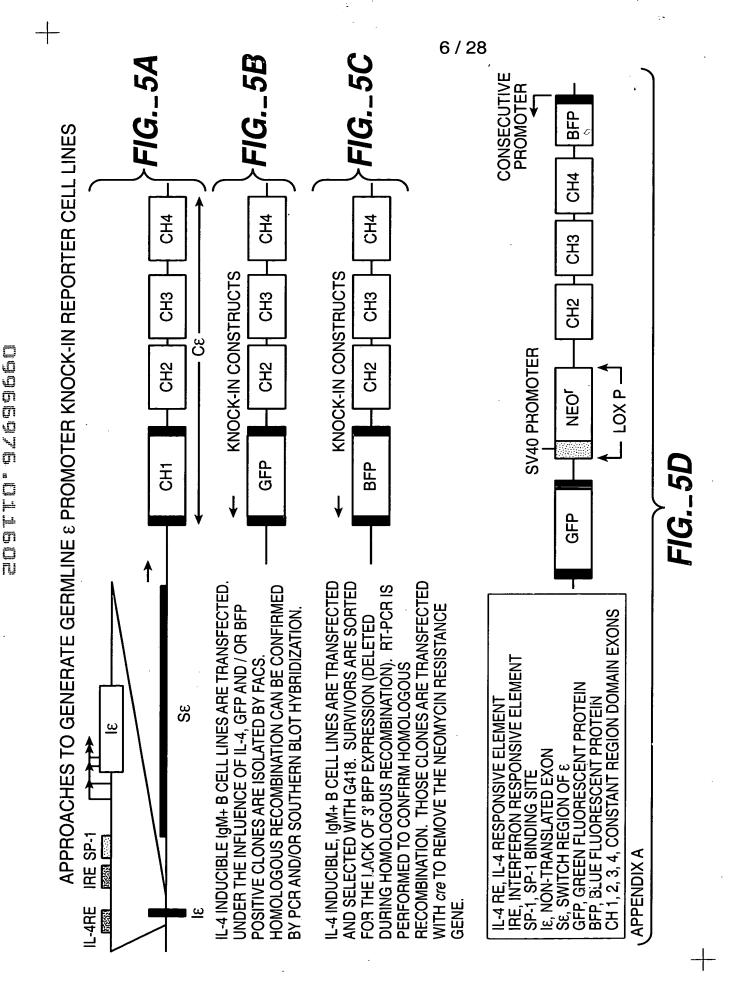
IL-4 INDUCTION OF GERMLINE ϵ mRNA IN THE IgM + B CELL LINES: CA-46, MC-116 AND DND39



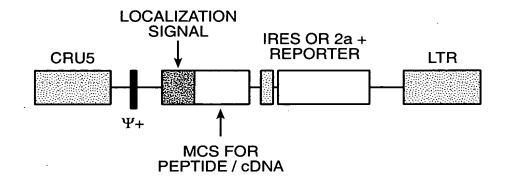
N

CELLS WERE INCUBATED FOR 48 HRS. IN 300 U / ml OF h-IL-4. RT-PCR WAS PERFORMED USING PRIMERS SPECIFIC FOR THE GERMLINE ϵ EXON AND THE 5'-END OF THE C ϵ CH1 EXON (PREDICTED SIZE \sim 200 bp). APPENDIX G

FIG._4



RIGEL BASE VECTOR



ALL COMPONENTS ARE UNIQUELY CASSETTED FOR FLEXIBILITY

CRU5, MODIFIED LTR
LTR, LONG TERMINAL REPEAT
Y+, PACKING SIGNAL
LOCALIZATION SIGNAL: NUCLEAR, CELL MEMBRANE, GRANULAR
MCS, MULTIPLE CLONING SITE
IRES, INTERNAL RIBOSOME ENTRY SITE
2a, SELF-CLEAVING PEPTIDE

APPENDIX I

FIG._6

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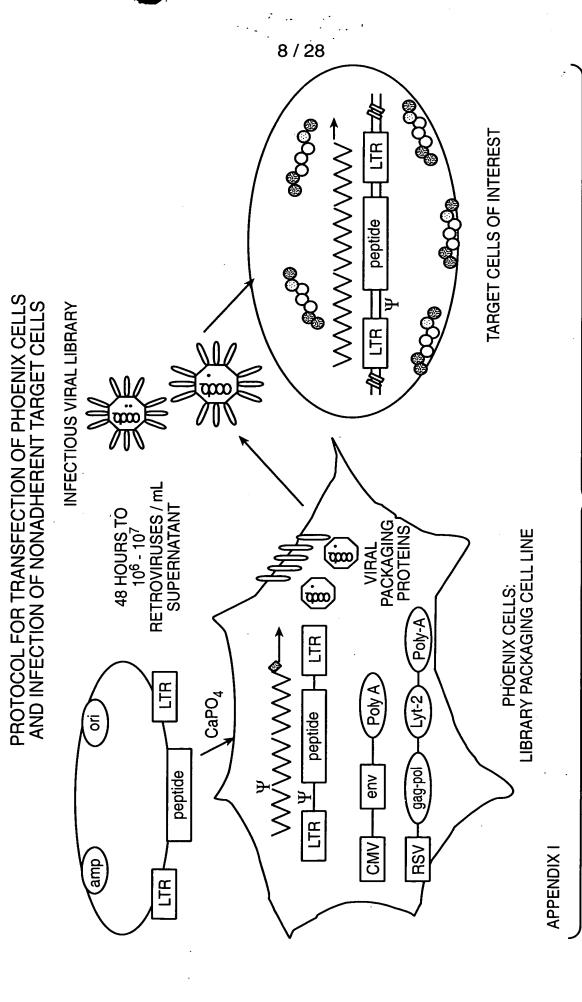


FIG._7

DOODONO OF HOOM

FIG._8B

HSV-TK, HERPES SIMPLEX VIRUS-THYMIDINE KINASE

CH 1, 2, 3, 4, CONSTANT REGION DOMAIN EXONS

M1, M2, MEMBRANE EXONS

Neo', NEOMYCIN RESISTANCE GENE BFP, BLUE FLUORESCENT PROTEIN

U266 CELLS ARE TRANSFECTED AND SELECTED WITH G418. SURVIVORS

HOMOLOGOUS RECOMBINATION). AT-PCR IS PERFORMED TO CONFIRM

ARE TREATED WITH GANCICLOVIR (HSV-TK DELETED DURING

HOMOLOGOUS RECOMBINATION. THOSE CLONES ARE TRANSFECTED

WITH cre TO REMOVE THE SV40 NEOMYCIN RESISTANCE GENE

APPENDIX D

VDJ, V REGION EXON

DOGENT DY PROPOS

FIG._9A

APPENDIX C

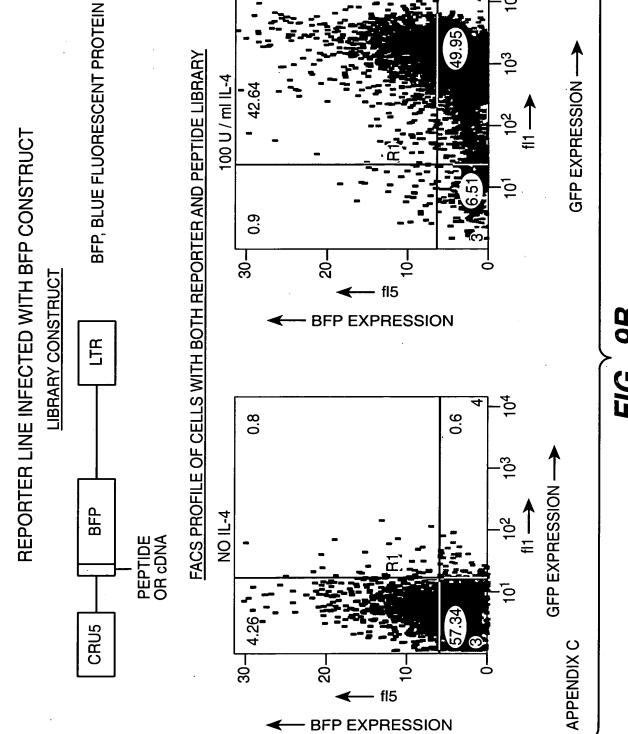
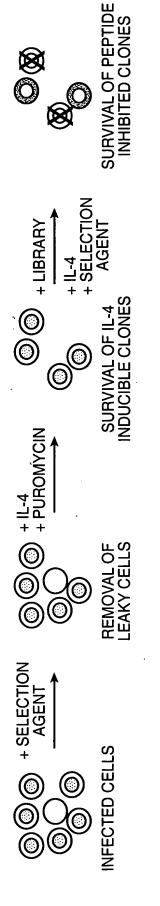


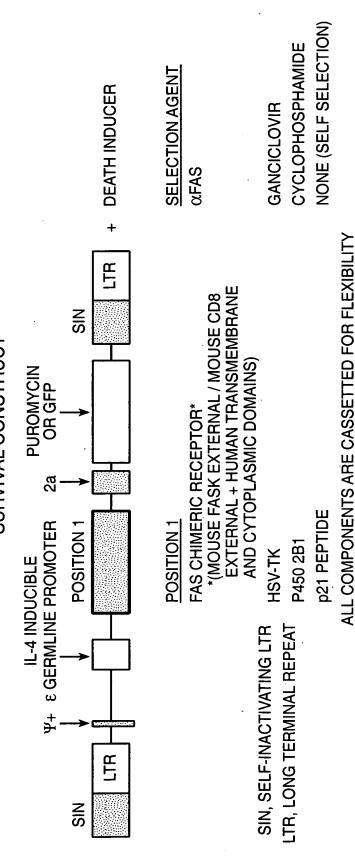
FIG._9B

NOUTIC " OZGODGGO

SCREEN FOR PEPTIDE INHIBITORS OF THE GERMLINE ϵ PROMOTER



SURVIVAL CONSTRUCT



APPENDIX D

FIG._ 10

1-845 CMV promoter/R/U5 5' LTR
1322 GAG ATG-ATC mutation
850-2100 extended Ψ region
2146-2173 two Bstx1 peptide cloning sites
2205-2723 ECMV IRES (cloned as EcoR1/Msc1 fragment from pCITE-4a [Novagen])
2746-3465 GFP coding region
3522-4115 3' LTR
4122-6210 pGEM backbone (pUC origin, ampR)

ATCACGAGGCCCTTTCGTCTTCAAGAACAGCTTTGCTCTTAGGAGTTTCCTAATACATCC CAAACTCAAATATAAAGCATTTGACTTGTTCTATGCCCTAGTTATTAATAGTAATCAA TTACGGGGTCATTAGTTCATAGCCCATATATGGAGTTCCGCGTTACATAACTTACGGTAA **ATGGCCCGCCTGGCTGACCGCCCAACGACCCCCCCCCATTGACGTCAATAATGACGTATG** TTCCCATAGTAACGCCAATAGGGACTTTCCATTGACGTCAATGGGTGGAGTATTTACGGT **AAACTGCCCACTTGGCAGTACATCAAGTGTATCATATGCCAAGTACGCCCCCTATTGACG** TCAATGACGGTAAATGGCCCGCCTGGCATTATGCCCAGTACATGACCTTATGGGACTTTC CTACTTGGCAGTACATCTACGTATTAGTCATCGCTATTACCATGGTGATGCGGTTTTTGGC **AGTACATCAATGGGCGTGGATAGCGGTTTGACTCACGGGGATTTCCAAGTCTCCACCCCA TTGACGTCAATGGGAGTTTGTTTTGGCACCAAAATCAACGGGACTTTCCAAAATGTCGTA** ACAACTCCGCCCCATTGACGCAAATGGGCGGTAGGCATGTACGGTGGGAGGTCTATATAA **GCAGAGCTCAATAAAAGAGCCCACAACCCCTCACTCGGGGCGCCAGTCCTCCGATTGACT** GAGTCGCCCGGGTACCCGTGTATCCAATAAACCCTCTTGCAGTTGCATCCGACTTGTGGT CTCGCTGTTCCTTGGGAGGGTCTCCTCTGAGTGATTGACTACCCGTCAGCGGGGGTCTTT CATTTGGGGGCTCGTCCGGGATCGGGAGACCCCTGCCCAGGGACCACCGACCACCG GGAGGTAAGCTGGCCAGCAACTTATCTGTGTCTGTCCGATTGTCTAGTGTCTATGACTGA TTTTATGCGCCTGCGTCGGTACTAGTTAGCTAACTAGCTCTGTATCTGGCGGACCCGTGG TGGAACTGACGAGTTCGGAACACCCGGCCGCAACCCTGGGAGACGTCCCAGGGACTTCGG **GGGCCGTTTTTGTGGCCCGACCTGAGTCCAAAAATCCCGATCGTTTTGGACTCTTTGGTG** CACCCCCTTAGAGGAGGGATATGTGGTTCTGGTAGGAGACGAGAACCTAAAACAGTTCC **GCTGCAGCATCGTTCTGTGTTGTCTCTGTCTGACTGTGTTTCTGTATTTGTCTGAAAATA** TCGGCCCGGGCCAGACTGTTACCACTCCCTTAAGTTTGACCTTAGGTCACTGGAAAGATG TCGAGCGGATCGCTCACAACCAGTCGGTAGATGTCAAGAAGAGACGTTGGGTTACCTTCT GCTCTGCAGAATGGCCAACCTTTAACGTCGGATGGCCGCGAGACGGCACCTTTAACCGAG **ACCTCATCACCCAGGTTAAGATCAAGGTCTTTTCACCTGGCCCGCATGGACACCCAGACC** CCTTTGTACACCCTAAGCCTCCGCCTCCTCTTCCTCCATCCGCCCCGTCTCTCCCCCTTG **AACCTCCTCGTTCGACCCCGCCTCGATCCTCCCTTTATCCAGCCCTCACTCCTTCTCTAG** GCGCCCCCATATGGCCATATGAGATCTTATATGGGGCACCCCCGCCCCTTGTAAACTTCC CTGACCCTGACATGACAAGAGTTACTAACAGCCCCTCTCTCCAAGCTCACTTACAGGCTC TCTACTTAGTCCAGCACGAAGTCTGGAGACCTCTGGCGGCAGCCTACCAAGAACAACTGG ACCGACCGGTGGTACCTCACCCTTACCGAGTCGGCGACACAGTGTGGGTCCGCCGACACC AGACTAAGAACCTAGAACCTCGCTGGAAAGGACCTTACACAGTCCTGCTGACCACCCCCA CCGCCCTCAAAGTAGACGGCATCGCGCTTGGATACACGCCGCCCACGTGAAGGCTGCCGA CCCCGGGGGTGGACCATCCTCTAGACTGCCGGATCTCGAGGGATCCACCACCATGGACCC

GGTTATTTTCCACCATATTGCCGTCTTTTGGCAATGTGAGGGCCCGGAAACCTGGCCCTG TCTTCTTGACGAGCATTCCTAGGGGTCTTTCCCCTCTCGCCAAAGGAATGCAAGGTCTGT TGAATGTCGTGAAGGAAGCAGTTCCTCTGGAAGCTTCTTGAAGACAACAACGTCTGTAG CGACCCTTTGCAGGCAGCGGAACCCCCCACCTGGCGACAGGTGCCTCTGCGGCCAAAAGC CACGTGTATAAGATACACCTGCAAAGGCGGCACAACCCCAGTGCCACGTTGTGAGTTGGA TAGTTGTGGAAAGAGTCAAATGGCTCTCCTCAAGCGTATTCAACAAGGGGCTGAAGGATG CCCAGAAGGTACCCCATTGTATGGGATCTGATCTGGGGCCTCGGTGCACATGCTTTACAT GTGTTTAGTCGAGGTTAAAAAACGTCTAGGCCCCCCGAACCACGGGGACGTGGTTTTCCT TTGAAAACACGATGATAATATGGGGGGATCCACCGGTCGCCACCATGGTGAGCAAGGGCG AGGAGCTGTTCACCGGGGTGGTGCCCATCCTGGTCGAGCTGGACGCGACGTAAACGGCC ACAAGTTCAGCGTGTCCGGCGAGGGCGAGGGCGATGCCACCTACGGCAAGCTGACCCTGA **AGTTCATCTGCACCACCGGCAAGCTGCCCGTGCCCTGGCCCACCCTCGTGACCACCCTGA** CCTACGGCGTGCAGTGCTTCAGCCGCTACCCCGACCACATGAAGCAGCACGACTTCTTCA **AGTCCGCCATGCCCGAAGGCTACGTCCAGGAGCGCACCATCTTCTTCAAGGACGACGGCA ACTACAAGACCCGCGCCGAGGTGAAGTTCGAGGGCGACACCCTGGTGAACCGCATCGAGC** TGAAGGGCATCGACTTCAAGGAGGACGGCAACATCCTGGGGCACAAGCTGGAGTACAACT **ACAACAGCCACAACGTCTATATCATGGCCGACAAGCAGAAGAACGGCATCAAGGTGAACT** TCAAGATCCGCCACAACATCGAGGACGGCAGCGTGCAGCTCGCCGACCACTACCAGCAGA ACACCCCATCGGCGACGGCCCGTGCTGCTGCCCGACAACCACTACCTGAGCACCCAGT CCGCCCTGAGCAAAGACCCCAACGAGAAGCGCGATCACATGGTCCTGCTGGAGTTCGTGA CCGCCGCCGGGATCACTCTCGGCATGGACGAGCTGTACAAGTAAAGCGGCCGCTCGACGA TAAAATAAAAGATTTTATTTAGTCTCCAGAAAAAGGGGGGGAATGAAAGACCCCACCTGTA **GGTTTGGCAAGCTAGCTTAAGTAACGCCATTTTGCAAGGCATGGAAAAATACATAACTGA** GAATAGAGAAGTTCAGATCAAGGTCAGGAACAGATGGAACAGCTGAATATGGGCCAAACA GGATATCTGTGGTAAGCAGTTCCTGCCCCGGCTCAGGGCCAAGAACAGATGGAACAGCTG **AATATGGGCCAAACAGGATATCTGTGGTAAGCAGTTCCTGCCCCGGCTCAGGGCCAAGAA** CAGATGGTCCCCAGATGCGGTCCAGCCCTCAGCAGTTTCTAGAGAACCATCAGATGTTTC CAGGGTGCCCCAAGGACCTGAAATGACCCTGTGCCTTATTTGAACTAACCAATCAGTTCG CTTCTCGCTTCTGTTCGCGCGCTTCTGCTCCCCGAGCTCAATAAAAGAGCCCACAACCCC TCACTCGGGGCGCCAGTCCTCCGATTGACTGAGTCGCCCGGGTACCCGTGTATCCAATAA **ACCCTCTTGCAGTTGCATCCGACTTGTGGTCTCGCTGTTCCTTGGGAGGGTCTCCTCTGA** GTGATTGACTACCCGTCAGCGGGGGTCTTTCATTTCCGACTTGTGGTCTCGCTGCCTTGG GAGGGTCTCCTCTGAGTGATTGACTACCCGTCAGCGGGGGTCTTCACATGCAGCATGTAT CAAAATTAATTTGGTTTTTTTTTCTTAAGTATTTACATTAAATGGCCATAGTTGCATTAAT GAATCGGCCAACGCGCGGGGAGAGGCGGTTTGCGTATTGGCGCTCTTCCGCTTCCTCGCT GGTAATACGGTTATCCACAGAATCAGGGGATAACGCAGGAAAGAACATGTGAGCAAAAGG CCAGCAAAAGGCCAGGAACCGTAAAAAGGCCGCGTTGCTGGCGTTTTTCCATAGGCTCCG CCCCCTGACGAGCATCACAAAAATCGACGCTCAAGTCAGAGGTGGCGAAACCCGACAGG **ACTATAAAGATACCAGGCGTTTCCCCCTGGAAGCTCCCTCGTGCGCTCTCCTGTTCCGAC** CCTGCCGCTTACCGGATACCTGTCCGCCTTTCTCCCTTCGGGAAGCGTGGCGCTTTCTCA TAGCTCACGCTGTAGGTATCTCAGTTCGGTGTAGGTCGTTCGCTCCAAGCTGGGCTGTGT GCACGAACCCCCGTTCAGCCCGACCGCTGCGCCTTATCCGGTAACTATCGTCTTGAGTC CAACCCGGTAAGACACGACTTATCGCCACTGGCAGCCACTGGTAACAGGATTAGCAG AGCGAGGTATGTAGGCGGTGCTACAGAGTTCTTGAAGTGGTGGCCTAACTACGGCTACAC

FIG._11A-2

TAGAAGGACAGTATTTGGTATCTGCGCTCTGCTGAAGCCAGTTACCTTCGGAAAAAGAGT **GCAGCAGATTACGCGCAGAAAAAAAGGATCTCAAGAAGATCCTTTGATCTTTCTACGGG GTCTGACGCTCAGTGGAACGAAAACTCACGTTAAGGGATTTTGGTCATGAGATTATCAAA** TATATATGAGTAAACTTGGTCTGACAGTTACCAATGCTTAATCAGTGAGGCACCTATCTC **AGCGATCTGTCTATTTCGTTCATCCATAGTTGCCTGACTCCCCGTCGTGTAGATAACTAC** GATACGGGAGGCTTACCATCTGGCCCCAGTGCTGCAATGATACCGCGAGACCCACGCTC ACCGGCTCCAGATTTATCAGCAATAAACCAGCCAGCCGGAAGGGCCGAGCGCAGAAGTGG TCCTGCAACTTTATCCGCCTCCATCCAGTCTATTAATTGTTGCCGGGAAGCTAGAGTAAG TAGTTCGCCAGTTAATAGTTTGCGCAACGTTGTTGCCATTGCTACAGGCATCGTGGTGTC ACGCTCGTCGTTTGGTATGGCTTCATTCAGCTCCGGTTCCCAACGATCAAGGCGAGTTAC **ATGATCCCCCATGTTGTGCAAAAAAGCGGTTAGCTCCTTCGGTCCTCCGATCGTTGTCAG AAGTAAGTTGGCCGCAGTGTTATCACTCATGGTTATGGCAGCACTGCATAATTCTCTTAC** TGTCATGCCATCCGTAAGATGCTTTTCTGTGACTGGTGAGTACTCAACCAAGTCATTCTG AGAATAGTGTATGCGGCGACCGAGTTGCTCTTGCCCGGCGTCAACACGCGGATAATACCGC GCCACATAGCAGAACTTTAAAAGTGCTCATCATTGGAAAACGTTCTTCGGGGCGAAAACT CTCAAGGATCTTACCGCTGTTGAGATCCAGTTCGATGTAACCCACTCGTGCACCCAACTG ATCTTCAGCATCTTTTACTTTCACCAGCGTTTCTGGGTGAGCAAAAACAGGAAGGCAAAA TCAATATTATTGAAGCATTTATCAGGGTTATTGTCTCATGAGCGGATACATATTTGAATG TATTTAGAAAAATAAACAAATAGGGGTTCCGCGCACATTTC

FIG._11A-3

1-845 CMVpromoter/R/U5 5' LTR
1322 GAG ATG-ATC mutation
850-2100 extend d ψ region
2151-2865 GFP coding region
2866-2894 GGGSGGG linker
2895-2952 FMDV 2a cleavage sequence
2953-3004 Bstx1/Bstx1/HinD3/Hpa1/Sal1/Not1 polylinker
3052-3645 3' LTR
3652-5715 pGEM backbone (pUC origin, ampR)

ATCACGAGGCCCTTTCGTCTTCAAGAACAGCTTTGCTCTTAGGAGTTTCCTAATACATC CCAAACTCAAATATATAAAGCATTTGACTTGTTCTATGCCCTAGTTATTAATAGTAATC **AATTACGGGGTCATTAGTTCATAGCCCATATATGGAGTTCCGCGTTACATAACTTACGG** TAAATGGCCCGCCTGGCTGACCGCCCAACGACCCCCGCCCATTGACGTCAATAATGACG TATGTTCCCATAGTAACGCCAATAGGGACTTTCCATTGACGTCAATGGGTGGAGTATTT **ACGGTAAACTGCCCACTTGGCAGTACATCAAGTGTATCATATGCCAAGTACGCCCCCTA** TTGACGTCAATGACGGTAAATGGCCCGCCTGGCATTATGCCCAGTACATGACCTTATGG GACTTTCCTACTTGGCAGTACATCTACGTATTAGTCATCGCTATTACCATGGTGATGCG **GTTTTGGCAGTACATCAATGGGCGTGGATAGCGGTTTGACTCACGGGGATTTCCAAGTC** TCCACCCCATTGACGTCAATGGGAGTTTGTTTTTGGCACCAAAATCAACGGGACTTTCCA **AAATGTCGTAACAACTCCGCCCCATTGACGCAAATGGGCGGTAGGCATGTACGGTGGGA GGTCTATATAAGCAGAGCTCAATAAAAGAGCCCACAACCCCTCACTCGGGGCGCCAGTC** CTCCGATTGACTGAGTCGCCCGGGTACCCGTGTATCCAATAAACCCTCTTGCAGTTGCA TCCGACTTGTGGTCTCGCTGTTCCTTGGGAGGGTCTCCTCTGAGTGATTGACTACCCGT CAGCGGGGGTCTTTCATTTGGGGGCTCGTCCGGGATCGGGAGACCCCTGCCCAGGGACC TAGTGTCTATGACTGATTTTATGCGCCTGCGTCGGTACTAGTTAGCTAACTAGCTCTGT **ATCTGGCGGACCCGTGGTGGAACTGACGAGTTCGGAACACCCGGCCGCAACCCTGGGAG ACGTCCCAGGGACTTCGGGGGCCGTTTTTGTGGCCCGACCTGAGTCCAAAAATCCCGAT** CGTTTTGGACTCTTTGGTGCACCCCCCTTAGAGGAGGATATGTGGTTCTGGTAGGAGA CGAGAACCTAAAACAGTTCCCGCCTCCGTCTGAATTTTTGCTTTCGGTTTGGGACCGAA TTTCTGTATTTGTCTGAAAATATCGGCCCGGGCCAGACTGTTACCACTCCCTTAAGTTT GACCTTAGGTCACTGGAAAGATGTCGAGCGGATCGCTCACAACCAGTCGGTAGATGTCA **AGAAGAGACGTTGGGTTACCTTCTGCTCTGCAGAATGGCCAACCTTTAACGTCGGATGG** CCGCGAGACGGCACCTTTAACCGAGACCTCATCACCCAGGTTAAGATCAAGGTCTTTTC **ACCTGGCCCGCATGGACACCCAGACCAGGTCCCCTACATCGTGACCTGGGAAGCCTTGG** CTTTTGACCCCCCTCCCTGGGTCAAGCCCTTTGTACACCCTAAGCCTCCGCCTCCTCTT CCTCCATCCGCCCGTCTCTCCCCCTTGAACCTCCTCGTTCGACCCCGCCTCGATCCTC CCTTTATCCAGCCCTCACTCCTTCTCTAGGCGCCCCCATATGGCCATATGAGATCTTAT **ATGGGGCACCCCGCCCCTTGTAAACTTCCCTGACCCTGACATGACAAGAGTTACTAAC AGCCCCTCTCTCCAAGCTCACTTACAGGCTCTCTACTTAGTCCAGCACGAAGTCTGGAG** GAGTCGGCGACACAGTGTGGGTCCGCCGACACCAGACTAAGAACCTAGAACCTCGCTGG **AAAGGACCTTACACAGTCCTGCTGACCACCCCCACCGCCCTCAAAGTAGACGGCATCGC** AGCTTGGATACACGCCGCCCACGTGAAGGCTGCCGACCCCGGGGGTGGACCATCCTCTA GACTGCCGGATCTCGAGGGATCCACCATGGTGAGCAAGGGCGAGGAGCTGTTCACCGGG

GTGGTGCCCATCCTGGTCGAGCTGGACGCGACGTAAACGGCCACAAGTTCAGCGTGTC CGGCGAGGGCGAGGCGATGCCACCTACGGCAAGCTGACCCTGAAGTTCATCTGCACCA CCGGCAAGCTGCCCGTGCCCTGGCCCACCCTCGTGACCACCCTGACCTACGGCGTGCAG TGCTTCAGCCGCTACCCCGACCACATGAAGCAGCACGACTTCTTCAAGTCCGCCATGCC CGAAGGCTACGTCCAGGAGCGCACCATCTTCTTCAAGGACGACGGCAACTACAAGACCC GCGCCGAGGTGAAGTTCGAGGGCGACACCCTGGTGAACCGCATCGAGCTGAAGGGCATC GACTTCAAGGAGGACGCCAACATCCTGGGGCACAAGCTGGAGTACAACTACAACAGCCA CAACGTCTATATCATGGCCGACAAGCAGAAGAACGGCATCAAGGTGAACTTCAAGATCC GCCACAACATCGAGGACGGCAGCGTGCAGCTCGCCGACCACTACCAGCAGAACACCCCC **ATCGGCGACGGCCCCGTGCTGCTGCCCGACAACCACTACCTGAGCACCCAGTCCGCCCT** GAGCAAAGACCCCAACGAGAAGCGCGATCACATGGTCCTGCTGGAGTTCGTGACCGCCG CCGGGATCACTCTCGGCATGGACGAGCTGTACAAGGAATTCGGAGGTGGCAGCGGTGGC GGTCAGCTGTTGAATTTTGACCTTCTTAAACTTGCGGGAGACGTCGAGTCCAACCCTGG GCCCACCACCACCATGGAAGCTTCCATTAAATTGGTTAACGTCGACGCGGCCGCTCGAC GATAAAATAAAAGATTTTATTTAGTCTCCAGAAAAAGGGGGGAATGAAAGACCCCACCT **GTAGGTTTGGCAAGCTAGCTTAAGTAACGCCATTTTGCAAGGCATGGAAAAATACATAA** CTGAGAATAGAGAAGTTCAGATCAAGGTCAGGAACAGATGGAACAGCTGAATATGGGCC **AAACAGGATATCTGTGGTAAGCAGTTCCTGCCCCGGCTCAGGGCCAAGAACAGATGGAA** CAGCTGAATATGGGCCAAACAGGATATCTGTGGTAAGCAGTTCCTGCCCCGGCTCAGGG CCAAGAACAGATGGTCCCCAGATGCGGTCCAGCCCTCAGCAGTTTCTAGAGAACCATCA GATGTTTCCAGGGTGCCCCAAGGACCTGAAATGACCCTGTGCCTTATTTGAACTAACCA **ATCAGTTCGCTTCTCGCTTCTGTTCGCGCGCTTCTGCTCCCCGAGCTCAATAAAAGAGC** CCACAACCCCTCACTCGGGGCGCCAGTCCTCCGATTGACTGAGTCGCCCGGGTACCCGT **GTATCCAATAAACCCTCTTGCAGTTGCATCCGACTTGTGGTCTCGCTGTTCCTTGGGAG** GGTCTCCTCTGAGTGATTGACTACCCGTCAGCGGGGGTCTTTCATTTCCGACTTGTGGT CTCGCTGCCTTGGGAGGGTCTCCTCTGAGTGATTGACTACCCGTCAGCGGGGGTCTTCA CATAGTTGCATTAATGAATCGGCCAACGCGCGGGGAGAGGCGGTTTGCGTATTGGCGCT CTTCCGCTTCCTCGCTCACTGACTCGCTGCGCTCGGTCGTTCGGCTGCGGCGAGCGGTA TCAGCTCACTCAAAGGCGGTAATACGGTTATCCACAGAATCAGGGGATAACGCAGGAAA GAACATGTGAGCAAAAGGCCAGCAAAAGGCCAGGAACCGTAAAAAAGGCCGCGTTGCTGG CGTTTTTCCATAGGCTCCGCCCCCTGACGAGCATCACAAAAATCGACGCTCAAGTCAG **AGGTGGCGAAACCCGACAGGACTATAAAGATACCAGGCGTTTCCCCCTGGAAGCTCCCT** CGTGCGCTCTCCTGTTCCGACCCTGCCGCTTACCGGATACCTGTCCGCCTTTCTCCCTT CGGGAAGCGTGGCGCTTTCTCATAGCTCACGCTGTAGGTATCTCAGTTCGGTGTAGGTC GTTCGCTCCAAGCTGGGCTGTGTGCACGAACCCCCCGTTCAGCCCGACCGCTGCGCCTT **ATCCGGTAACTATCGTCTTGAGTCCAACCCGGTAAGACACGACTTATCGCCACTGGCAG** CAGCCACTGGTAACAGGATTAGCAGAGCGAGGTATGTAGGCGGTGCTACAGAGTTCTTG **AAGTGGTGGCCTAACTACGGCTACACTAGAAGGACAGTATTTGGTATCTGCGCTCTGCT** CAAGAAGATCCTTTGATCTTTTCTACGGGGTCTGACGCTCAGTGGAACGAAAACTCACG TTAAGGGATTTTGGTCATGAGATTATCAAAAAGGATCTTCACCTAGATCCTTTTAAATT **AAAAATGAAGTTTGCGCAAATCAATCTAAAGTATATATGAGTAAAACTTGGTCTGACAGT** TACCAATGCTTAATCAGTGAGGCACCTATCTCAGCGATCTGTCTATTTCGTTCATCCAT **AGTTGCCTGACTCCCCGTCGTGTAGATAACTACGATACGGGAGGGCTTACCATCTGGCC** CCAGTGCTGCAATGATACCGCGAGACCCACGCTCACCGGCTCCAGATTTATCAGCAATA

FIG._11B-2

FIG._11B-3

1-845 CMVpormoter/R/U5 5' LTR
1322 GAG ATG-ATC mutation
850-2100 extended ψ region
2146-2173 two Bstx1 peptide cloning sites
2173-2214 EoR1/Apa1/Hpa1/Not1 polylinker
2262-2855 3' LTR
2855-4901 pGEM backbone (pUC origin, ampR)

ATCACGAGGCCCTTTCGTCTTCAAGAACAGCTTTGCTCTTAGGAGTTTCCTAATACATC CCAAACTCAAATATAAAGCATTTGACTTGTTCTATGCCCTAGTTATTAATAGTAATC **AATTACGGGGTCATTAGTTCATAGCCATATATGGAGTTCCGCGTTACATAACTTACGGT AAATGGCCCGCCTGGCTGACCGCCCAACGACCCCCGCCCATTGACGTCAATAATGACGT ATGTTCCCATAGTAACGCCAATAGGGACTTTCCATTGACGTCAATGGGTGGAGTATTTA** CGGTAAACTGCCCACTTGGCAGTACATCAAGTGTATCATATGCCAAGTACGCCCCCTAT TGACGTCAATGACGGTAAATGGCCCGCCTGGCATTATGCCCAGTACATGACCTTATGGG **ACTITICATE TAGGE AGAINATION ACTIVITY OF A** TTTTGGCAGTACATCAATGGGCGTGGATAGCGGTTTGACTCACGGGGATTTCCAAGTCT CCACCCCATTGACGTCAATGGGAGTTTGTTTTGGCACCAAAATCAACGGGACTTTCCAA **AATGTCGTAACAACTCCGCCCCATTGACGCAAATGGGCGGTAGGCATGTACGGTGGGAG GTCTATATAAGCAGAGCTCAATAAAAGAGCCCACAACCCCTCACTCGGGGCGCCAGTCC** TCCGATTGACTGAGTCGCCCGGGTACCCGTGTATCCAATAAACCCTCTTGCAGTTGCAT CCGACTTGTGGTCTCGCTGTTCCTTGGGAGGGTCTCCTCTGAGTGATTGACTACCCGTC AGCGGGGGTCTTTCATTTGGGGGCTCGTCCGGGATCGGGAGACCCCTGCCCAGGGACCA **AGTGTCTATGACTGATTTTATGCGCCTGCGTCGGTACTAGTTAGCTAACTAGCTCTGTA** TCTGGCGGACCCGTGGTGGAACTGACGAGTTCGGAACACCCGGCCGCAACCCTGGGAGA CGTCCCAGGGACTTCGGGGGCCGTTTTTGTGGCCCGACCTGAGTCCAAAAATCCCGATC **GTTTTGGACTCTTTGGTGCACCCCCTTAGAGGAGGGATATGTGGTTCTGGTAGGAGAC** GAGAACCTAAAACAGTTCCCGCCTCCGTCTGAATTTTTGCTTTCGGTTTGGGACCGAAG TTCTGTATTTGTCTGAAAATATCGGCCCGGGCCAGACTGTTACCACTCCCTTAAGTTTG **ACCTTAGGTCACTGGAAAGATGTCGAGCGGATCGCTCACAACCAGTCGGTAGATGTCAA** GAAGAGACGTTGGGTTACCTTCTGCTCTGCAGAATGGCCAACCTTTAACGTCGGATGGC CGCGAGACGGCACCTTTAACCGAGACCTCATCACCCAGGTTAAGATCAAGGTCTTTTCA CCTGGCCCGCATGGACACCCAGACCAGGTCCCCTACATCGTGACCTGGGAAGCCTTGGC TTTTGACCCCCCTCCCTGGGTCAAGCCCTTTGTACACCCTAAGCCTCCGCCTCCTCTTC CTCCATCCGCCCCGTCTCTCCCCCTTGAACCTCCTCGTTCGACCCCCGCCTCGATCCTCC CTTTATCCAGCCCTCACTCCTTCTCTAGGCGCCCCCATATGGCCATATGAGATCTTATA TGGGGCACCCCCGCCCCTTGTAAACTTCCCTGACCCTGACATGACAAGAGTTACTAACA GCCCCTCTCTCCAAGCTCACTTACAGGCTCTCTACTTAGTCCAGCACGAAGTCTGGAGA CCTCTGGCGGCAGCCTACCAAGAACAACTGGACCGACCGGTGGTACCTCACCCTTACCG **AGTCGGCGACACAGTGTGGGTCCGCCGACACCAGACTAAGAACCTAGAACCTCGCTGGA AAGGACCTTACACAGTCCTGCTGACCACCCCCACCGCCTCAAAGTAGACGGCATCGCA** GCTTGGATACACGCCGCCCACGTGAAGGCTGCCGACCCCGGGGGTGGACCATCCTCTAG **ACTGCCGGATCTCGAGGGATCCACCACCATGGACCCCCATTAAATTGGAATTCGGGGCC** TAACGCCATTTTGCAAGGCATGGAAAAATACATAACTGAGAATAGAGAAGTTCAGATCA

AGGTCAGGAACAGATGGAACAGCTGAATATGGGCCAAACAGGATATCTGTGGTAAGCAG TTCCTGCCCCGGCTCAGGGCCAAGAACAGATGGAACAGCTGAATATGGGCCAAACAGGA TATCTGTGGTAAGCAGTTCCTGCCCCGGCTCAGGGCCAAGAACAGATGGTCCCCAGATG CGGTCCAGCCCTCAGCAGTTTCTAGAGAACCATCAGATGTTTCCAGGGTGCCCCAAGGA CCTGAAATGACCCTGTGCCTTATTTGAACTAACCAATCAGTTCGCTTCTCGCTTCTGTT CGCGCGCTTCTGCTCCCCGAGCTCAATAAAAGAGCCCCACAACCCCTCACTCGGGGCGCC AGTCCTCCGATTGACTGAGTCGCCCGGGTACCCGTGTATCCAATAAACCCTCTTGCAGT TGCATCCGACTTGTGGTCTCGCTGTTCCTTGGGAGGGTCTCCTCTGAGTGATTGACTAC CCGTCAGCGGGGGTCTTTCATTTCCGACTTGTGGTCTCGCTGCCTTGGGAGGGTCTCCT **CTGAGTGATTGACTACCCGTCAGCGGGGGTCTTCACATGCAGCATGTATCAAAATTAAT** TTGGTTTTTTTTCTTAAGTATTTACATTAAATGGCCATAGTTGCATTAATGAATCGGCC **AACGCGCGGGGAGAGGCGGTTTGCGTATTGGCGCTCTTCCGCTTCCTCGCTCACTGACT** CGGTTATCCACAGAATCAGGGGATAACGCAGGAAAGAACATGTGAGCAAAAGGCCAGCA AAAGGCCAGGAACCGTAAAAAGGCCGCGTTGCTGGCGTTTTTCCATAGGCTCCGCCCCC CTGACGAGCATCACAAAAATCGACGCTCAAGTCAGAGGTGGCGAAACCCGACAGGACTA TAAAGATACCAGGCGTTTCCCCCTGGAAGCTCCCTCGTGCGCTCTCCTGTTCCGACCCT GCCGCTTACCGGATACCTGTCCGCCTTTCTCCCTTCGGGAAGCGTGGCGCTTTCTCATA GCTCACGCTGTAGGTATCTCAGTTCGGTGTAGGTCGTTCGCTCCAAGCTGGGCTGTGTG CACGAACCCCCGTTCAGCCCGACCGCTGCGCCTTATCCGGTAACTATCGTCTTGAGTC CAACCGGTAAGACACGACTTATCGCCACTGGCAGCAGCCACTGGTAACAGGATTAGCA GAGCGAGGTATGTAGGCGGTGCTACAGAGTTCTTGAAGTGGTGGCCTAACTACGGCTAC **ACTAGAAGGACAGTATTTGGTATCTGCGCTCTGCTGAAGCCAGTTACCTTCGGAAAAAG AGTTGGTAGCTCTTGATCCGGCAAACAAACCACCGCTGGTAGCGGTGGTTTTTTTGTTT** GCAAGCAGCAGATTACGCGCAGAAAAAAAGGATCTCAAGAAGATCCTTTGATCTTTTCT ACGGGGTCTGACGCTCAGTGGAACGAAAACTCACGTTAAGGGATTTTGGTCATGAGATT **ATCAAAAAGGATCTTCACCTAGATCCTTTTAAATTAAAAATGAAGTTTGCGCAAATCAA** TCTAAAGTATATATGAGTAAACTTGGTCTGACAGTTACCAATGCTTAATCAGTGAGGCA CCTATCTCAGCGATCTGTCTATTTCGTTCATCCATAGTTGCCTGACTCCCCGTCGTGTA GATAACTACGATACGGGAGGGCTTACCATCTGGCCCCAGTGCTGCAATGATACCGCGAG CGCAGAAGTGGTCCTGCAACTTTATCCGCCTCCATCCAGTCTATTAATTGTTGCCGGGA **AGCTAGAGTAAGTTCGCCAGTTAATAGTTTGCGCAACGTTGTTGCCATTGCTACAG** TCAAGGCGAGTTACATGATCCCCCATGTTGTGCAAAAAAGCGGTTAGCTCCTTCGGTCC TCCGATCGTTGTCAGAAGTAAGTTGGCCGCAGTGTTATCACTCATGGTTATGGCAGCAC TGCATAATTCTCTTACTGTCATGCCATCCGTAAGATGCTTTTCTGTGACTGGTGAGTAC TCAACCAAGTCATTCTGAGAATAGTGTATGCGGCGACCGAGTTGCTCTTGCCCGGCGTC **AACACGGGATAATACCGCGCCACATAGCAGAACTTTAAAAAGTGCTCATCATTGGAAAAC GTTCTTCGGGGCGAAAACTCTCAAGGATCTTACCGCTGTTGAGATCCAGTTCGATGTAA** CCCACTCGTGCACCCAACTGATCTTCAGCATCTTTTACTTTCACCAGCGTTTCTGGGTG **AGCAAAAACAGGAAGGCAAAATGCCGCAAAAAAGGGAATAAGGGCGACACGGAAATGTT** GAATACTCATACTCTTTCTTTTCAATATTATTGAAGCATTTATCAGGGTTATTGTCTC ATGACATTAACCTATAAAAATAGGCGT

FIG._11C-2

(1) C12ScFas Survival construct

psilon-cFas(CD95)-Ires-Hygro-BGH PolyA put into C12s vector backwards so that no leaky transcription happens through the cmv promoter.

ttttatgegeetgegteggtaetagttagetaaetagetetgtatetggeggaeeegtggtggaaetgaegagtteggaa tegttttggaetetttggtgeaceeeettagaggagggatatgtggttetggtaggagaegagaacetaaaaeagttee tgtctctgtctgactgtgtttctgtatttgtctgaaaatatgggcccgggccagactgttaccactcccttaagtttgac gctctgcagaatggccaacctttaacgtcggatggccgcgagacggcacctttaaccgagacctcatcacccaggttaag atcaaggtetttteacetggeeegeatggaeaceeagaeeaggteeeetaeategtgaeetgggaageettggettttga ccccctccctgggtcaagccctttgtacaccctaagcctccgcctcctcttcctccatccgccccgtctcccccttg accgaccggtggtacctcacccttaccgagtcggcgacacagtgtgggtccgccgacaccagactaagaacctagaacct cgctggaaaggaccttacacagtcctgctgaccaccccaccgccctcaaagtagacggcatcgcagcttggatacacgc cgcccacgtgaaggctgccgaccccggggggtggaccatcctctagactgccGGATCTCGAGGATCCTCCCCAGCATGCC atcacgaggccctttcgtcttcaagaacagctttgctcttaggagtttcctaatacatcccaaactcaaatatataaagc atttgacttgttctatgccctagttattaatagtaatcaattacggggtcattagttcatagcccatatatggagttccg cgttacataacttacggtaaatggcccgcctggctgaccgcccaacgacccccgcccattgacgtcaataatgacgtatg ttoccatagtaacgocaatagggactttocattgacgtcaatgggtggagtatttacggtaaactgcccacttggcagta catcaagtgtatcatatgccaagtacgccccctattgacgtcaatgacggtaaatggcccgcctggcattatgcccagta catgaccttatgggactttcctacttggcagtacatctacgtattagtcatcgctattaccatggtgatgcggtttttggc agtacatcaatgggcgtggatagcggtttgactcacggggatttccaagtctccaccccattgacgtcaatgggagtttg ttttggcaccaaaatcaacgggactttccaaaatgtcgtaacaactccgccccattgacgcaaatgggcggtaggcatgt acggtgggaggtctatataagcagagctcaataaaagagcccacacaaccctcactcgggggggcgccagtcctccgattgact gagtegeeegggtaceegtgtateeaataaaeeetettgeagttgeateegaettgtggtetegetgtteettggggaggg totoototgagtgattgactacccgtcagcgggggtctttcatttgggggctcgtccggggatcgggagacccctgcccag ggaccaccaccaccgggaggtaagctggccagcaacttatctgtgtccgattcgtctagtgtctagtgtctatgactga caccoggcogcaaccctgggagacgtcccagggacttcgggggccgtttttgtgggcccgacctgagtccaaaaatcccga cttaggtcactggaaagatgtcgagcggatcgctcacaaccagtcggtagatgtcaagaagagacgttgggttaccttct ccaagctcacttacaggctctctacttagtccagcacgaagtctggagacctctggcggcagcctaccaagaacaactgg

 gcaaacaacagatggctggcaactagaaggcacagtcgaggtctagcttgccaaacctacaggtggggtcttcattcc $FIG_{-1}2A$

FIG._ 12B

TTTTTACCAGGTTGGCATGGTTGACAGCAAAATGGGCCTCCTTGATAAATCCTTCTGAGCAGTTTTTATCAGTTTTCATG

gegacetegtattgggaateecegaacategeetegeteeagteaatgacegetgttatgeggeeattgteegteaggae attgttggagccgaaatccgcgtgcacgaggtgccggacttcggggcagtcctcggcccaaagcatcagctcatcgagag cetgogogacggacgcactgacggtgtcgtccatcacagttttgccagtgatacacatggggatcagcaatcgcgcatatg aaatcacgccatgtagtgtattgaccgattccttgcggtccgaatgggccgaacccgctcgtctggctaagatcggccgc cctgtgcacggcgggagatgcaataggtcaggctctcgctaaattccccaatgtcaagcacttccggaatcgggagcgcg googatgoaaagtgoogataaacataacgatotttgtagaaaccatoggogoagotatttaocogoaggacatatocaog ccetcctacatcgaagctgaaagcacgagattettcgccetccgagagetgcatcaggtcggagacgetgtcgaaetttt cgatcagaaacttctcgacagacgtcgcggtgagttcaggctttttcatggtattatcatcgtgtttttcaaaggaaaac cacgtccccgtggttccgggggcctagacgtttttaacctcgactaaacatgtaaagcatgtgcaccgaggccccag atcagatcccatacaatggggtaccttctgggcatccttcagccccttgttgaatacgcttgaggagagccatttgactc tttccacaactatccaactcacaacgtggcactggggttgtgccgcctttgcaggtgtatcttatacacgtggcttttgg ccaagctttggatttcatttctgaagtttgaattttctgagtcactagtaatgtccttgaggatgatgatgtctgaattttc ccaattacgaagcagttgaactttctgttctgctgtcttggacattgtcattcttgatctcatctcatttttggcttcat tgacaccattctttcgaacaaagcctttaacttgacttagtgtcatgactccagcaatagtggtgatatatttactcaag tcaacatcagataaatttattgccactgtttcaggatttaaggttggagattcatgagaaccttggttttcctttctgtg cttctgcatgttttctgtacttccttctctcacccaaacaattagtggaattggcaaaagaagacaaagccacc ccaaccggTTTCTGGGACTTTGTTTCCTGCAGTTTGTATTGCTGGTTGCTGTGCATGGCTCAAGGGTTCCATGTTCACAC **L** L C C A G A G C A C C T T C A C A T T C A C A C C T T G C A T T C C T T T G G C G A G G G G A A G A C C C L A G A C L A G <u>GAGGCGCAGCGAACACACAGTGTTCACAGCCAGGAGAATCGCAGTAGAAGTCTGGTTTGCACTTGCACTTGGTATTCTGGGT</u> CAGGGTGCAGTTTGTTTCCACTTCTAAACCATGCTCTTCATCGCAGAGTGTGCATCTTCTGCATTTATCAGCATAATGGT

DOSESTE STABLE

gegatttgtgtaegecegaeagteeeggeteeggateggaegattgegtegeategaecetgegeeeaagetgeateate gaaattgccgtcaaccaagctctgatagagttggtcaagaccaatgcggagcatatacgcccggagccgcgggggatcctg caageteeeggatgeeteegetegaagtagegegtetgetgeteeaataeaageeaaeeaeggeeteeagaagatgttg

CCCTTTTTCTGGAGACTAAATAAATCTTTTATTTTatcgatagatcccggtcggcatctactctattcctttgccctcg gacgagtgctggggcgtcggtttccactatcggcgagtacttctacacagccatcggtccagacggccgcgcttctgcgg

gregegaaaccegacagactataaagataccagecettrececetegaagetecetegtegegetecegateegacee FIG._12C CtagcTTAAGTAACCCATTTTGCAAGGCATGGAAAAATACATAACTGAGAATAGAGAAGGTCAGATCAAGGTCGGAACAG **ATGGAACAGGCAATAAAAGAGCCCACAACCCCTCACTCGGGGCGCCAGTCCTCCGATTGACTGAGTCGCCGGGTACCCG GGCCATagtttcGTAATCATGGTCATAGCTGTTTCCTGTGAAATTGTTATCCGCTCACAATTCCACAACATACGAG TTCCAGTCGGGAAACCTGTCGTGCCAGCTGCATTAATGAATCGGCCAACGCGCGGGGAGAGAGGCGGTTTGCGTATTGGGCG AAACGCAAGAGTCTTCTCTGTCTCGACAAGCCCAGTTTCTATTGGTCTCCTTAAACCTGTCTTGTAACCTTGATACTTAC** gtgccgggcagtggagcctgggtagggggagctctgcctcagtgctttcagctaaaaatgggggtgggaaccccCaggagg cccgggccgccctggaagttcccttttctctctgttcttgggaagtcgattgagcaacagcgggggtcaggtgaggctcc ttcactaccgatgcacaccgagtgctGggggggggttctctctctctcaggcccaacCccagggcccctgcctaggtccc ggactetCactettgaegeatgegtggettggtggteceagteageaaattggggteeegttgeetgggaaagggagag cgcactgaggtgaactggccctcggggGcgcgtgtcccagatgtgtgtgcagggcctcctgatggccgcagcctcgtcc ctgtgacccgcttggagctggcacctgagtggcctcacCTTGTACTCCCCAGGTCACTGTCTcgacGCGGCC GCTCGAcgataaaataaaagattttatttagtctccagaaaaaggggggaatgaaagaccccacctgtaggtttggcaag TGTATCCAATAAACCCTCTTGCAGTTGCATCCGACTTGTGGTCTCGCTGTTCCTTGGGAGGGTCTCCTCTGAGTGTGTTGA CTACCCGTCAGCGGGGGTCTTTCAcatgcaGCATGTATCAAAATTTAATTTGGTTTTTTTTTTTTAAGTATTTACATTAAAT TAATACGGTTATCCACAGAATCAGGGGATAACGCAGGAAAGAACATGTGAGCAAAAAGGCCAGCAAAAGGCCAGGAAACGT aaaaaggccgcgttgcttgcgttttttcataggctccgcccctgacgagcatcaaaaaatcgacgcctcaagag CTGCCCAGTGCCTCACGACCAACTTctgcaggaattcctggacagctcccagatgatcagtaaccgtggttgttatttct

DOOLLO OLOGOOLO

CCAGAGGCAGGACAGCCCAGATCCACACCATgGTGGCTTTACCAACAGTACCGGAATGCCAAGCTTGCGGCCGCTTAAGA

GCTGTAATTGAACCTGGGAGTGGACACCTGTGGAGAGAAAGGCAAAGTGGATGTCAGTAAGACCAATAGGTGCCTATCAG

TAACTATCGTCTTGAGTCCAACCCGGTAAGACACGACTTATCGCCACTGGCAGCAGCACTGGTAACAGGATTAGCAGGA CCGGGAAGCTAGAGTAAGTTCGCCCAGTTAATAGTTTGCGCAACGTTGTTGCCATTGCTACAGGCATCGTGGTGTCACA **ACTGCATAATTCTCTTTACTGTCATGCCATCCGTAAGATGCTTTTCTGTGACTGGTGagtactcaaccaagtcattctgag** aatagtgtatgcggcgaccgagttgctcttgcccggcgtcaacacgggataataccgcgccacatagcagaactttaaaa TGCCGCTTACCGGATACCTGTCCGCCTTTCTCCCTTCGGGAAGCGTGGCGCTTTCTCATAGCTCACGCTGTAGGTATCTC <u> AGTTCGGTGTAGGTCGTTCGCTCCAAGCTGGGCTGTGTGCACGAACCCCCCGTTCAGCCCGACCGCTGCGCCTTATCCGG</u> **CGAGGTATGTAGGCGGTGCTACAGAGTTCTTGAAGTGGTGGCCTAACTACGGCTACACTAGAAGGACAGTATTTGGTATC** TACGGGAGGGCTTACCATCTGGCCCCCAGTGCTGCAATGATACCGCGAGACCCCACGCTCACCGGCTCCAGATTTATCAGCA GCTCGTCGTTTGGTATGGCTTCATTCAGCTCCGGTTCCCAACGATCAAGGCGAGTTACATGATCCCCCATGTTGTGCAAA CTGACGCTCAGTGGAACGAAAACTCACGTTAAGGGATTTTGGTCATGAGATTATCAAAAAGGATCTTCACCTAGATCCTT

MODITIO, WYGDDGGO

FIG._ 12D

gtgeteateattggaaaacgttetteggggegaaaaeteteaaggatettaeegetgttgagateeggttegatgtaaee cactegtgeacecaactgatetteageatettttaettteaceagegttetgggtgageaaaaaggaaggeaaatg

cogcaaaaaagggaataagggogacacggaaatgttgaatactcatactcttccttttcaatattattgaagcatttat

cagggttattgtctcatgacattaacctataaaaataggcgt

caageteeggatgeeteegetegaagtagegegtetgetgeteeataeaageeaaceaeggeeteeagaagatgttg gogacotogtattgggaatcocogaacatcgcotogotocagtcaatgacogotgttatgoggcoattgtocgtcaggac attgttggagccgaaatccgcgtgcacgaggtgccggacttcggggcagtcctcggcccaaagcatcagctcatcgagag cotgogogacgacgactgacggtgtcgtccatcacagtttgccagtgatacacatggggatcagcaatcgcgtatatg aaatcacgccatgtagtgtattgaccgattccttgcggtccgaatgggccgaacccgctcgtctggctaagatcggccgc cctgtgcacggcgggagatgcaataggtcaggctctcgctaaattccccaatgtcaagcacttccggaatcgggagcgcg gccgatgcaaagtgccgataaacataacgatctttgtagaaaccatcggcgcagctatttacccgcaggacatatccacg coctoctacatogaagctgaaagcacgagattottcgccctccgagagctgcatcaggtcggagacgctgtcgaactttt cgatcagaaacttctcgacagacgtcgcgggtgagttcaggctttttcatggtattatcatcgtgttttttcaaaggaaaac cacgtccccgtggttccggggggcctagacgtttttaacctcgactaaacatgtaaagcatgtgcaccgaggccccag atcagatcccatacaatggggtaccttctgggcatccttcagccccttgttgaatacgcttgaggaggccatttgactc tttccacaactatccaactcacaacgtggcactggggttgtgcgcctttgcaggtgtatcttatacacgtggcttttgg **LtCCAGAGGAACTGCTTCCTTCACGACATTCAACAGACCTTGCATTCCTTTGGCGAGAGGGGAAAGACCCCtagactaga**

tgacaccattotttcgaacaaagcotttaacttgacttagtgtcatgactccagcaatagtggtgatatatttactcaag cttctgcatgttttctgtacttcctttctcttcacccaaacaattagtggaattggcaaaagaagacaaagccacc tgcacaggtgagggagttcgcagcactggcttggtagtagtagaggtcacttctgaaggactggcacgacagaactgaa gtacatcaccgagttgctgatgactgagcagaaatagtagccttcgttttccttgctgaacttgttcagggtgagaacgt gaagccatatagacaacgaaggtgggctggggggggttttggagctggagttctggaagagccaagagcatccttgcgaaac gtgcctgtggcttagcttctccactccccaggataatcgactcacccagcagcagcaggtgctcagcgacagaaagcgggtc ccaagctttggatttcatttctgaagtttgaattttctgagtcactagtaatgtccttgaggatgatagtctgaatttttc ccaattacgaagcagttgaactttctgttctgctgtgttcttggacattgtcattcttgatctcatctattttggcttcat tcaacatcagataaatttattgccactgtttcaggatttaaggttggagattcatgagaaccttggttttcctttctgtg ccaaccggtttccggtccccttcactgagccacggggccgacaatcttctggtctctggggctgagatgtcccggtaggg acttattattegtgteeeteatggeagaaaacagtttegaegaatteagettetegteeegttatettgttgtgggat ggaccccaacacttcacataccaggtccaccttctgaccaagttcggcgtccattttctttggaaagattcggagttcgg aacggtgaggccatgGTGGCTTTACCAACAGTACCGGAATGCCAAGCTTGCGGCCGCTTAAGAGCTGTAATTGAACCTGG

ctgectegacaageceagtetetategetetetetaaacetgetetegaacetegatatetatetageceagtgecteacg $F\!\!\!/\!\!\!/\!\!\!/ 5B$ GAGTGGACACCTGTGGAGAAAAGGCAAAGTGGATGTCAGTAAGACCAATAGGTGCCTATCAGAAACGCAAGAGTCTTCT

gacgagtgctgggggcgtcggtttccactatcggcgagtacttctacacagccatcggtccagacggccgcgcttctgcgg gaaattgocgtcaaccaagottotgatagagttggtcaagaccaatgoggagcatatacgocoggagooggoggogatootg

CCCTTTTTCTGGAGACTAAATAAATCTTTTATTTTA toga taga to cogg togg ca to tactotatto ot ttg coctog

- (2) Ahhh: Survival construct
- 2.) Abbbh: epsilon-cFas' (CD8 or mLyt2)-Ires-Hygro-BGHpolyA also in C12s backwards

caccoggcogcaaccotgggaagacgtcocagggacttcgggggcogtttttgtggcocgacctgagtccaaaaatcocga tegttttggaetetttggtgeaceeeettagaggagggatatgtggttetggtaggagaegagaacetaaaaeagttee tgtctctgtctgactgtgtttctgtatttgtctgaaaatatgggcccgggccagactgttaccactcccttaagtttgac gctctgcagaatggccaacctttaacgtcggatggccgcgagacggcacctttaaccgagacctcatcacccaggttaag atcaaggtetttteaeetggeeegeatggaeaeeeeagaeeaggteeeetaeategtgaeetgggaageettggettttga coccettoctgggtcaagccetttgtacaccetaagcetccgcetcetettectccatccgccccgtctcctccccttg atttgacttgttctatgccctagttattaatagtaatcaattacggggtcattagttcatagcccatatatggagttccg cgttacataacttacggtaaatggcccgcctggctgaccgcccaacgacccccgcccattgacgtcaataatgacgtatg ttoccatagtaacgocaatagggactttocattgacgtcaatgggtggagtatttacggtaaactgcccacttggcagta catcaagtgtatcatatgccaagtacgccccctattgacgtcaatgacggtaaatggcccgcctggcattatgcccagta catgaccttatgggactttcctacttggcagtacatctacgtattagtcatcgctattaccatggtgatgcggtttttggc agtacatcaatgggcgtggatagcggtttgactcacggggatttccaagtctccaccccattgacgtcaatgggagtttg tttggcaccaaaatcaacgggactttccaaaatgtcgtaacaactccgccccattgacgcaaatgggcggtaggcatgt gagtegeeegggtaceegtgtateeaataaaeeetettgeagttgeateegaettgtggtettgetgetgtteettggggaggg tctcctctgagtgattgactacccgtcagcgggggtctttcatttggggggctcgtccggggatcgggagacccctgcccag tttatgegeetgegteggtaetagttagetaactagetetgtatetggeggegegegggggggggaaetgaeggtggtggaa cttaggtcactggaaagatgtcgagcggatcgctcacaaccagtcggtagatgtcaagaagaggcggttgggttaccttct aacctcctcgttcgaccccgcctcgatcctccctttatccagccctccactccttctctaggcgcccccatatggccatat gagatettatatggggeaceeegeeettgtaaaetteeetgaeeetgaeeatgaeaagagttaetaaeageeeetetet ccaagetcaettacaggetetetaettagtecageacgaagtetggagaeetetggeggeggeageetaeeaagaaeaaetgg accgaccggtggtacctcacccttaccgagtcggcgacacagtgtgggtccgccgacaccagactaagaacctagaacct cgctggaaaggaccttacacagtcctgctgaccacccccaccgccttcaaagtagacggcatcgcagcttggatacacgc cgcccacgtgaaggctgccgaccccggggggtggaccatcctagactgccGGATCTCGAGGGATCTCCTCCCCAGCATGCC TGCTATTGTCTTCCCAATCCTCCCCCTTGCTGTCCTGCCCCACCCCCACCCCCAGAATAGAATGACACCTACTCAGACAA atcacgaggccctttcgtcttcaagaacagctttgctcttaggagtttcctaatacatcccaaactcaaatatataaagc

gcaaacaacagatggctggcaactagaaggcacagtcgaggtctagcttgccaaacctacaggtggggtcttcattcc $\,FIG_-13A$

CTTGCAGTTGCATCCGACTTGTGGTCTCGCTGTTCCTTGGGAGGGTCTCCTCTGAGTGATTGACTACCCGTCAGCGGGG

TCTTTCAcatgcaGCATGTATCAAAATTAATTTGGTTTTTTTTTTTTAAGTATTTAAATTGGCCATAGttcGTAAT

CATGGTCATAGCTGTTTCCTGTGTGAAATTGTTATCCGCTCACAATTCCACACATACGAGCCGGAAGCATAAAGTGT

GTCGTGCCAGCTGCATTAATGAATCGGCCAACGCGGGGAGAGGGGGGTTTGCGTATTGGGCGCTCTTCCGCTTCCTCGC

AAAGCCTGGGGTGCCTAATGAGTGAGCTAACTCACATTAATTGCGTTGCGCTCACTGCCCGCTTTCCAGTCGGGAAACCT

CCAACCCGGTAAGACACGACTTATCGCCACTGGCAGCACTGGTAACAGGATTAGCAGAGCGAGGTATGTAGGCGGT FIG. 13C GGCGTTTTTCCATAGGCTCCGCCCCCTGACGAGCATCACAAAAATCGACGCTCAAGTCAGAGGTGGCGAAACCCGACAG GACTATAAAGATACCAGGCGTTTCCCCCTGGAAGCTCCCTCGTGCGCTCTCTTCCGACCCTGCCGCTTACCGGATAC CTGTCCGCCTTTCTCCCTTCGGGAAGCGTGGCGCTTTCTCATAGCTCACGCTGTAGGTATCTCAGTTCGGTGTAGGTCGT Gaatcaggggataacgcaggaaagaacatgtgagcaaaaggccagcaaaaaggccaggaaccgtaaaaaggccgcgttgct TCGCTCCAAGCTGGGCTGTGTGCACGAACCCCCCGTTCAGCCCGACCGCTGCGCCTTATCCGGTAACTATCGTCTTGAGT

DODHHO" DYGODGOO

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TTTTGCAAGGCATGGAAAAATACATAACTGAGAATAGAGAAGTTCAGATCAAGGTCGGAACAGATGGAACAGGCAATAAA

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GAAAACTCACGTTAAGGGATTTTGGTCATGAGATTATCAAAAAGGATCTTCACCTAGATCCTTTTAAATTAAAATGAAG GTAGTTCGCCAGTTAATAGTTTGCGCAACGTTGTTGCCATTGCTACAGGCATCGTGGTGTCACGCTCGTCGTTTGGTATG gatottcagcatottttactttcaccagcgtttctgggtgagcaaaaacaggaaggcaaaatgccgcaaaaagggaata GCTACAGAGTTCTTGAAGTGGTGGCCTAACTACGGCTACACTAGAAGGACAGTATTTGGTATCTGCGCTCTGCTGAAGCC **AGCAGCAGATTACGCGCAGAAAAAAAGGATCTCAAGAAGATCCTTTGATCTTTTCTACGGGGTCTGACGCTCAGTGGAAC** TTTGCGCAAATCAATCTAAAGTATATATGAGTAAACTTGGTCTGACAGTTACCAATGCTTAATCAGTGAGGCACCTATCT CAGCGATCTGTCTATTTCGTTCATCCATAGTTGCCTGACTCCCGTCGTGTAGATAACTACGATACGGGAGGGCTTACCA GCTTCATTCAGCTCCGGTTCCCAACGATCAAGGCGAGTTACATGATCCCCCCATGTTGTGCAAAAAAGCGGTTAGCTCCTT CGGTCCTCCGATCGTTGTCAGAAGTAAGTTGGCCGCAGTGTTATCACTCATGGTTATGGCAGCACTGCATAATTCTCTTA CTGTCATGCCATCCGTAAGATGCTTTTCTGTGACTGGTGagtactcaaccaagtcattctgagaatagtgtatgcggcga ccgagttgctcttgcccggcgtcaacacggggataataccgcgccacatagcagaactttaaaagtgctcatcattggaaa acgttettegggggggaaaaeteteaaggatettacegetgttgagateeagttegatgtaaeeeaetegtgeaeeeaaet agggcgacacggaaatgttgaatactcatactcttcctttttcaatattattgaagcatttatcagggttattgtctcat gacattaacctataaaaataggcgt

FIG._ 13D